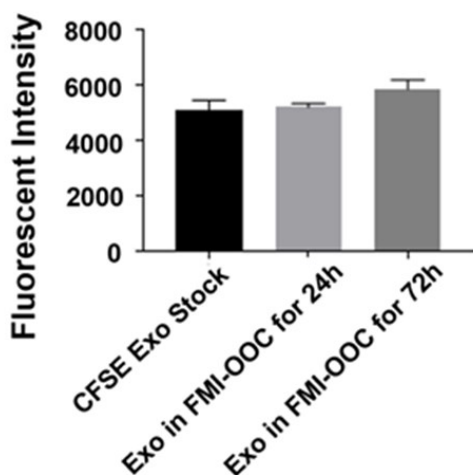
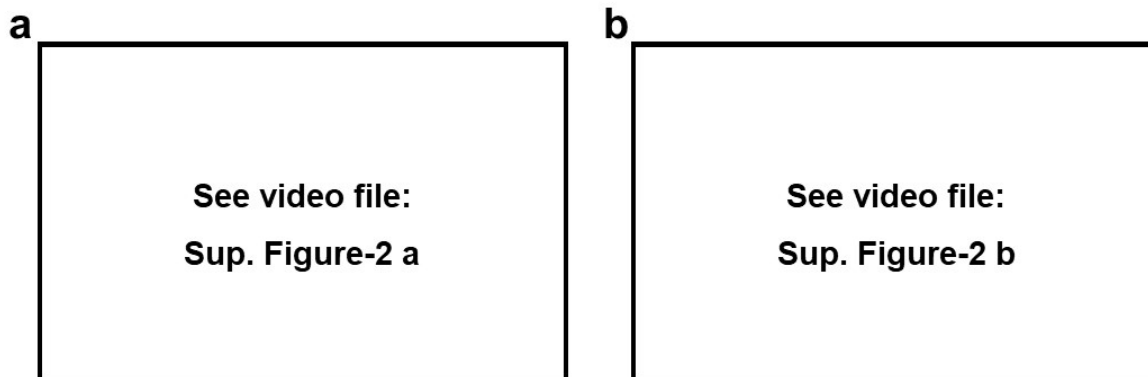


SUPPLEMENTAL FIGURES



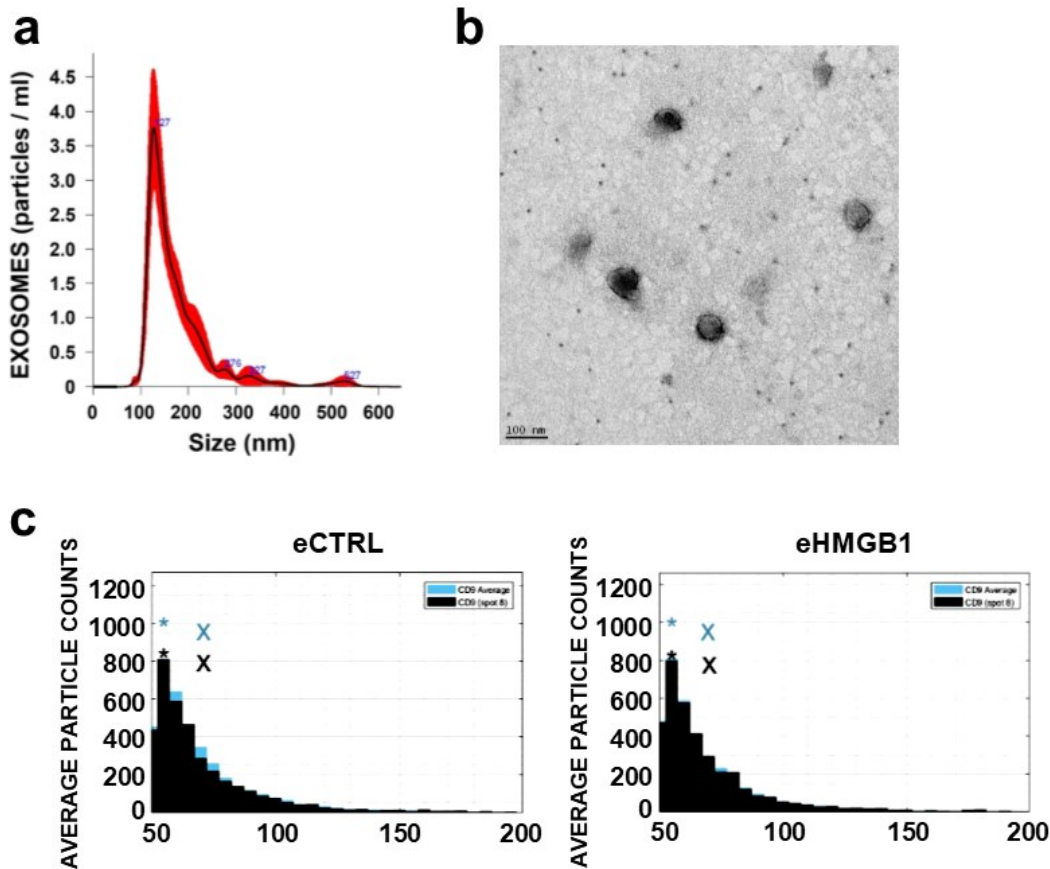
Supplementary Fig.1 Fluorescently labeled exosomes trafficking through the layers of cell lines in FMI-OOC device.

Fluorescent plate reader measured CFSE-labeled fluorescent exosome concentrations within the FMI-OOC, showing no difference over 72 h period, indicating no or limited absorption of exosomes to the PDMS device or reservoir system ($n = 3$). Values are expressed as means \pm SEM.



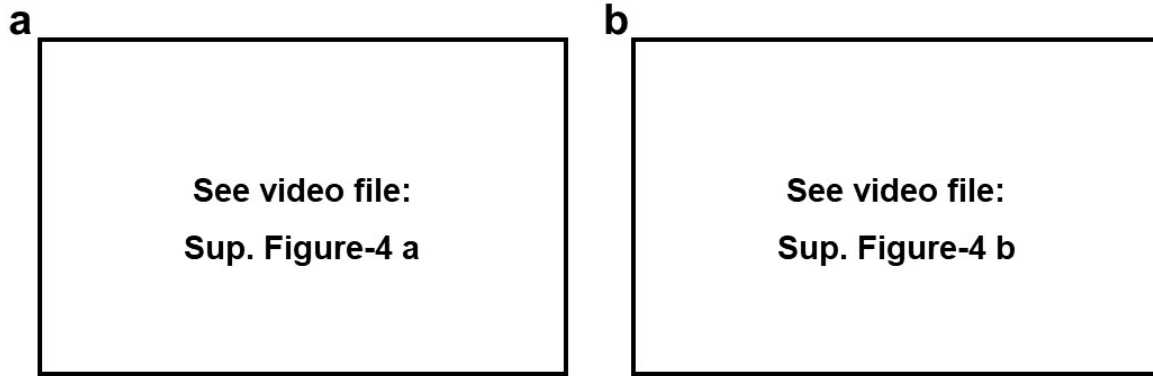
Supplementary Fig.2 Time-lapse movie of transiently expressed HMGB1-GFP colocalization with CD9-RFP in amnion epithelial cells (CD9-RFP-AECs) under oxidative stress condition. **(a)** Transiently expressed HMGB1-GFP translocation into the cytoplasm

against oxidative stress (OS) induction by cigarette smoke extract (CSE) within 24 h. **(b)** Colocalization of HMGB1-GFP with CD9-RFP in AECs induced by OS. Time-lapse captured for 24 h with Keyence microscope.

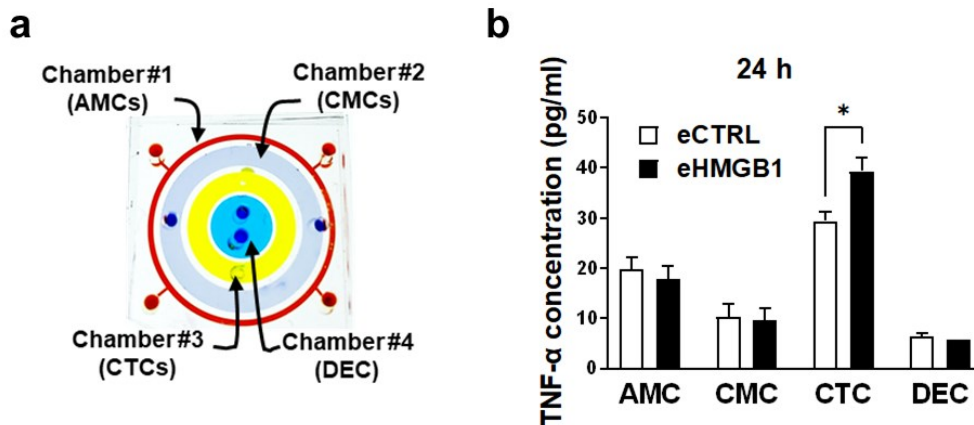


Supplementary Fig.3 Characterization of AEC-derived exosomes. **(a)** Size distribution of AEC-derived exosomes before electroporation with nanoparticle tracking analysis (NTA) (n=5). **(b)**. Representative wide-field transmission electron microscopy (TEM) images of AEC-derived exosomes before electroporation (N=3). **(c)** Representative graphs of engineered eCTRL and eHMGB1 CD9-captured exosomes

size distribution by ExoView analysis.

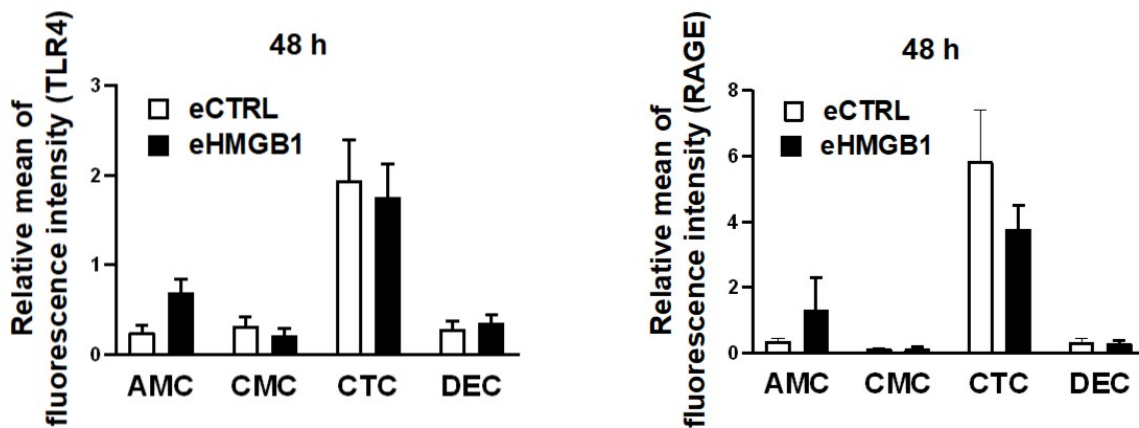


Supplementary Fig.4 Exosomes trafficking through the layers of cell lines in FMI-OOC device. **(a)** Time-lapse movies of endogenous and exogenous exosomes trafficking across cell layers. **(b)** movie merged with phase contrast. Scale Bars, 50 μ m.

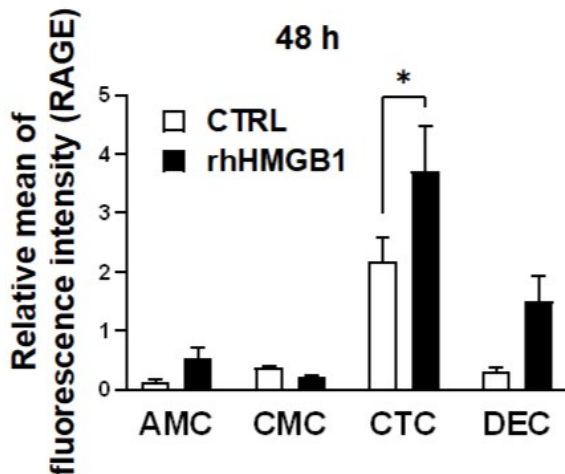


Supplementary Fig.S5 Engineered HMGB1-exosomes trafficking initiates the activation of inflammatory cytokine production throughout fetal membrane cells

(a) Image of the FMI-OOC for inflammation propagation study. AMCs (chamber 1), CMCs (chamber 2), CTCs (chamber 3) and DEC (chamber 4) were seeded as indicated. **(b)** Secreted cytokine TNF- α level was measured by immunoassay using the culture media from 24 h treatment (n=3). The data is presented as the means \pm SEM. *P < 0.05, two-way analysis of variance (ANOVA).



Supplementary Fig.S6 Engineered HMGB1-exosomes trafficking is blocked no channel FMI-OOC devices. Fluorescence signal intensity of TLR4 and RAGE expression was measured at indicated time periods by ImageJ (n=3). The data is presented as the means \pm SEM. two-way analysis of variance (ANOVA).



Supplementary Fig.S7 rhHMGB1 treatment induced RAGE expression in FMI-OOC devices. Fluorescence signal intensity of RAGE expression was measured at 48 h by ImageJ (n=3). The data is presented as the means \pm SEM. $P=0.016$, two-way analysis of variance (ANOVA).